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NEWS	4	JAN 28	USPATFULL, USPAT2, and USPATOLD enhanced with new custom IPC display formats
NEWS	5	JAN 28	MARPAT searching enhanced
NEWS	6	JAN 28	USGENE now provides USPTO sequence data within 3 days of publication
NEWS	7	JAN 28	TOXCENTER enhanced with reloaded MEDLINE segment
NEWS	8	JAN 28	MEDLINE and LMEDLINE reloaded with enhancements
NEWS	9	FEB 08	STN Express, Version 8.3, now available
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NEWS	11	FEB 25	IFIREF reloaded with enhancements
NEWS	12	FEB 25	IMSPRODUCT reloaded with enhancements
NEWS	13	FEB 29	WPINDEX/WPIDS/WPIX enhanced with ECLA and current U.S. National Patent Classification
NEWS	14	MAR 31	IFICDB, IFIPAT, and IFIUDB enhanced with new custom IPC display formats
NEWS	15	MAR 31	CAS REGISTRY enhanced with additional experimental spectra
NEWS	16	MAR 31	CA/CAPLUS and CASREACT patent number format for U.S. applications updated
NEWS	17	MAR 31	LPCI now available as a replacement to LDPCI
NEWS	18	MAR 31	EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS	19	APR 04	STN AnaVist, Version 1, to be discontinued
NEWS	20	APR 15	WPIDS, WPINDEX, and WPIX enhanced with new predefined hit display formats
NEWS	21	APR 28	EMBASE Controlled Term thesaurus enhanced
NEWS	22	APR 28	IMSRESEARCH reloaded with enhancements
NEWS EXPRESS	FEBRUARY 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 20 FEBRUARY 2008		
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NEWS IPC8	For general information regarding STN implementation of IPC 8		

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\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 10:07:43 ON 19 MAY 2008

=> file caplus medline biosis embase

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

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0.21

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FILE 'EMBASE' ENTERED AT 10:08:08 ON 19 MAY 2008

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=> s biotin

L1 95250 BIOTIN

=> s benzene

L2 399378 BENZENE

=> s aspartyl

L3 18645 ASPARTYL

=> s L1 and L2

L4 173 L1 AND L2

=> s L3 and L4

L5 0 L3 AND L4

=> s ethers or carboxylates or sulfonates or ammonium

L6 811867 ETHERS OR CARBOXYLATES OR SULFONATES OR AMMONIUM

=> s L4 and L6

L7 9 L4 AND L6

=> dup rem L7

PROCESSING COMPLETED FOR L7

L8 9 DUP REM L7 (0 DUPLICATES REMOVED)

=> d 1-9 L8 ibib abs

L8 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:485496 CAPLUS

DOCUMENT NUMBER: 146:462680

TITLE: Water-soluble electropolymerizable monomers having metalloporphyrin groups

INVENTOR(S): Canonne, Frederic; Korri, Youssoufi Hafsa; Mahy, Jean Pierre; Mandrand, Bernard; Perree Fauvet, Martine

PATENT ASSIGNEE(S): Biomerieux, Fr.; Centre National de la Recherche Scientifique; Universite Paris Sud (Paris XI)

SOURCE: Fr. Demande, 57pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent

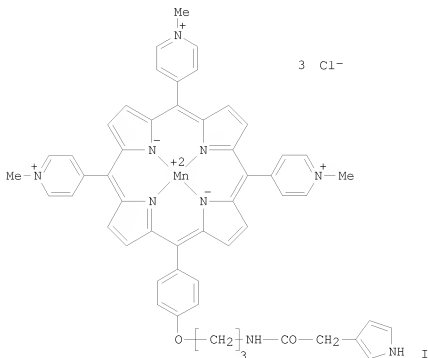
LANGUAGE: French  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR 2892723	A1	20070504	FR 2005-11187	20051103
WO 2007051947	A1	20070510	WO 2006-FR51131	20061102
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				

PRIORITY APPLN. INFO.: FR 2005-11187 A 20051103

OTHER SOURCE(S): MARPAT 146:462680

GI



AB Metalloporphyrins are manufactured having electropolymerizable groups and  $\geq 2$  groups that are ionizable or ionized in water. Thus, reaction of 4-hydroxybenzaldehyde with N-(3-bromopropyl)phthalimide, cyclization of the intermediate with 4-pyridinecarboxaldehyde and pyrrole, hydrolysis of the resulting porphyrin derivative, reaction of the resulting porphyrin derivative having a 3-aminopropoxy group with 1H-pyrrol-3-ylacetic acid, methylation

of the resulting porphyrin derivative having an amide group, and metalation of the resulting porphyrin deriv salt with Mn gave a complex I, which exhibited electropolymerizability.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:1270480 CAPLUS

DOCUMENT NUMBER: 147:507931

TITLE: Procedure for the biodegradation of MTBE AND TBA in the presence of additional organic pollutants by means of carrier-fixed microorganisms

INVENTOR(S): Rohwerder, Thore; Mueller, Roland H.; Martienssen, Marion

PATENT ASSIGNEE(S): Ufz-Umweltforschungszentrum Leipzig-Halle G.m.b.H., Germany

SOURCE: Ger. Offen., 8pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 102006022042	A1	20071108	DE 2006-102006022042	20060505
PRIORITY APPLN. INFO.: DE 2006-102006022042 20060505				
AB The invention concerns a procedure for the biol. degradation of Me tert. Bu ether (MTBE) and/or tert-Bu alc. (TBA), optionally in presence of further pollutants from contaminated waters. In addition to MTBE and TBA, BTEX compds. (benzene, toluene, ethylbenzene, xylene) as well as petroleum mineral oil hydrocarbons can be simultaneously eliminated. The invention concerns a bacterial strain specialized for the complete mineralization of MTBE, Ideonella sp. L-108, , which is settled on a substrate, used for the biodegrdn. of the contaminant(s). The characteristics of the carrier are specifically optimized for the settlement of these bacteria. According to the invention substrates with a moderately hydrophobic character, without or with weakly cationic net charge, are preferable, including burned clay granulate, lava tuff and polystyrene.				

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 3 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2006:907178 CAPLUS

DOCUMENT NUMBER: 146:337875

TITLE: A process for the preparation of substituted 6-benzyl-5-oxo-3-phenyl-(3S,7S,7aR)-perhydroimidazo[1,5-c][1,3]thiazoles

INVENTOR(S): Chavan, Subhash Prataprao; Gopal, Chittiboyina Amar; Kamat, Subhash Krishnaji; Kalkote, Uttam Ramrao; Ravindranathan, Thotapallil

PATENT ASSIGNEE(S): Council of Scientific and Industrial Research, India

SOURCE: Indian, 17pp.

CODEN: INXXAP

DOCUMENT TYPE: Patent

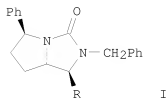
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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IN 194355	A1	20041023	IN 2001-DE302	20010319
US 6486328	B1	20021126	US 2000-686908	20001012
PRIORITY APPLN. INFO.:			US 2000-686908	A 20001012
OTHER SOURCE(S):	CASREACT 146:337875; MARPAT 146:337875			
GI				



AB A process for the preparation of substituted 6-benzyl-5-oxo-3-phenyl-(3S,7S,7aR)-perhydroimidazo[1,5-c][1,3]thiazoles having general formula I [R = 1-phenyl-1-ethanone, 1-(4-chlorophenyl)-1-ethanone, 1-(4-methoxyphenyl)-1-ethanone, 2-oxocyclohexyl, 1-trimethylsilyloxy-2-oxocyclohexyl, allyl, 1-hexynyl, 4-dimethyl aminophenyl, or 2-methylpropanoate] which comprises reacting the compound 6-benzyl-7-hydroxy-3-phenyl-(3S,7aR)-perhydroimidazo[1,5-c][1,3]thiazol-5-one as described herein with Lewis acid and a nucleophile at a temperature ranging between 0 to 30°C in an organic solvent for 10-30 min, quenching the reaction mixture with quenching agent (preferably with water and/or saturated aqueous solution of salts of sodium, potassium, ammonium), separating and concentrating the organic layer, and purifying by known conventional methods such as chromatog. (silica gel) to obtain I. I have the potential as serving as intermediates to biotin or as chiral auxiliaries.

L8 ANSWER 4 OF 9 CAPLUS COPYRIGHT 2008 ACS on SIN

ACCESSION NUMBER: 2004:525055 CAPLUS  
DOCUMENT NUMBER: 141:76350  
TITLE: Oxidative hair dyes containing derivatives of indoline, p-phenylene diamine and conditioners  
INVENTOR(S): Kleen, Astrid  
PATENT ASSIGNEE(S): Henkel Kommanditgesellschaft Auf Aktien, Germany  
SOURCE: Eur. Pat. Appl., 24 pp.  
CODEN: EPXXDW  
DOCUMENT TYPE: Patent  
LANGUAGE: German  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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EP 1433470	A1	20040630	EP 2003-29369	20031219
EP 1433470	B1	20080326		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
DE 10260835	A1	20040701	DE 2002-10260835	20021223
AT 390118	T	20080415	AT 2003-29369	20031219
PRIORITY APPLN. INFO.:			DE 2002-10260835	A 20021223
OTHER SOURCE(S):	MARPAT 141:76350			

AB The invention concerns oxidative hair dyes that contain in a cosmetically acceptable carrier (a) at least one indoline derivative; (b) at least one p-phenylene diamine derivative; (c) hair care substances selected from the group of (c1) vitamins and provitamins, e.g. vitamins A, B, E and H; (c2) ectoin and its derivs., allantoin and bisabolol. Addnl. developers and cationic direct dyes can be added. Thus a hair dye cream contained

(weight/weight%): Texapon NSO 15; Dehyton K 12; Lorol 4; Hydrenol 10; Eumulgin B1 0.5; Eumulgin B2 0.5; ascorbic acid 0.5; sodium sulfite 0.5; ammonium hydrogenphosphate 1; ammonia to pH10; 1-(2-hydroxyethyl)-2,5-diaminobenzene sulfate 1.3; 5,6-dihydroxyindoline hydrobromide 0.1; 2-methylresorcin 0.09; 4-chlororesorcin 0.3; n-aminophenol 0.05; 2,7-dihydroxynaphthalene 0.23; 2,4,5,6-tetraaminopyrimidine 0.03; 1-methoxy-2-amino-4-(2-hydroxyethylamino) benzene sulfate 0.03; Vitamin B6 0.5; water to 100.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2002:503386 CAPLUS

DOCUMENT NUMBER: 137:63120

TITLE: Process for preparing perhydroimidazol[1,5-c][1,3]thiazol-5-one derivatives as intermediates for synthesis of D(+)-biotin

INVENTOR(S): Chavan, Subhash Prataprao; Chittiboyina, Amar Gopal; Kamat, Subhash Krishnaji; Kalkote, Uttam Ramrao; Ravindranathan, Thotapallil

PATENT ASSIGNEE(S): Council of Scientific and Industrial Research, India

SOURCE: Eur. Pat. Appl., 14 pp.

CODEN: EPXXDW

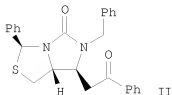
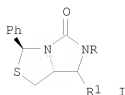
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1219625	A1	20020703	EP 2000-128571	20001227
EP 1219625	B1	20060517		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
AT 326471	T	20060615	AT 2000-128571	20001227
PT 1219625	T	20061031	PT 2000-128571	20001227
ES 2264919	T3	20070201	ES 2000-128571	20001227
PRIORITY APPLN. INFO.:			EP 2000-128571	A 20001227
OTHER SOURCE(S):	CASREACT 137:63120; MARPAT 137:63120			
GI				



AB The present invention discloses a process for preparing substituted perhydroimidazol[1,5-c][1,3]thiazol compds., such as I [R = benzyl; R1 = alkyl group exemplified by 1-phenyl-1-ethanone, 1-(4-chlorophenyl)-1-ethanone,  $\alpha$ -(4-methoxyphenyl)-1-ethanone, 2-oxocyclohexyl, 1-trimethylsilyloxy-2-oxocyclohexyl, allyl, 1-hexanyl, etc.], as crucial intermediates for the synthesis of D(+)-biotin. Thus, D(+)-biotin intermediate II was prepared by the reaction of 1-trimethylsilyloxystyrene and 6-benzyl-7-hydroxy-3-phenyl-(3S,7aR)-perhydroimidazol[1,5-c][1,3]thiazol-5-one in presence of boron trifluoride etherate. These compds. are more stable and are produced by non-hazardous

methods.  
REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2008 ACS on SIN  
ACCESSION NUMBER: 2000:861452 CAPLUS  
DOCUMENT NUMBER: 134:29252  
TITLE: Synthesis of water soluble multi-biotin  
-containing compounds for use in targeting  
biotin-binding proteins  
PATENT ASSIGNEE(S): University of Washington, USA  
SOURCE: PCT Int. Appl., 68 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 5  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000072802	A2	20001207	WO 2000-US15081	20000601
WO 2000072802	A3	20020207		
W: AU, BR, CA, IL, JP, KR, MX, RU				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 1196199	A2	20020417	EP 2000-938025	20000601
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				

PRIORITY APPLN. INFO.: US 1999-324267 A 19990602  
WO 2000-US15081 W 20000601

AB Syntheses of water soluble discrete multi-biotin-containing compds.  
with at least three biotin moieties are disclosed. The water  
soluble biotin-containing compds. may addnl. comprise one or more  
moieties that confer resistance to cleavage by biotinidase or that is  
cleavable in vitro or in vivo. The discrete multi-biotin-containing  
compds. may include a reactive moiety that provides a site for reaction  
with yet another moiety, such as a targeting, diagnostic or therapeutic  
functional moiety. Biotinylation reagents comprising water soluble linker  
moieties are also disclosed and may addnl. comprise a biotinidase  
protective group. Methods for amplifying the number of sites for binding  
biotin-binding proteins at a selected target using multi-  
biotin compds. are also disclosed.

L8 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2008 ACS on SIN  
ACCESSION NUMBER: 1999:784331 CAPLUS  
DOCUMENT NUMBER: 132:20747  
TITLE: Surface regeneration of biosensors using a combination  
of solutions based on interaction-specific optimized  
processes  
INVENTOR(S): Andersson, Karl; Hamalainen, Markku; Malmqvist,  
Magnus; Roos, Hakan  
PATENT ASSIGNEE(S): Biacore AB, Swed.  
SOURCE: PCT Int. Appl., 133 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9963333	A1	19991209	WO 1999-SE921	19990531

W: AU, JP, US  
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,  
PT, SE

US 6289286	B1	20010911	US 1998-87402	19980529
AU 9946658	A	19991220	AU 1999-46658	19990531
AU 755181	B2	20021205		
EP 1082607	A1	20010314	EP 1999-930044	19990531

R: BE, CH, DE, FR, GB, LI, NL, SE, FI

JP 2002517720	T	20020618	JP 2000-552490	19990531
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PRIORITY APPLN. INFO.: US 1998-87402 A 19980529

WO 1999-SE921 W 19990531

AB Surface regeneration of affinity biosensors and characterization of biomols. associated therewith by multivariate technique employing cocktails of regeneration agents to optimize regeneration of biosensor surface and/or characterize biomols. associated therewith. Kits and stock solns. for use in the context of this invention, as well as associated computer algorithms are also disclosed. Stock solns. of regeneration cocktails are prepared and combined. Solns. are acidic, basic, ionic, organic, detergent and chelating agent containing Biosensors for various affinity bindings are regenerated by the method; the affinity reactions are used for optimizing the regeneration process. Immuno-reactions, nucleic acid hybridization, avidin/streptavidin-biotin, hormone-hormone receptor interactions are performed with Biocore instruments and CM5 sensor chips.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1966:11410 CAPLUS

DOCUMENT NUMBER: 64:11410

ORIGINAL REFERENCE NO.: 64:2057h,2058a-b

TITLE: Substituted indole derivatives

INVENTOR(S): Shavel, John, Jr.; Strandtmann, Maximilian Von

PATENT ASSIGNEE(S): Warner-Lambert Pharmaceutical Co.

SOURCE: 2 pp.

DOCUMENT TYPE: Patent

LANGUAGE: Unavailable

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	---	-----	-----	-----
US 3217029		19651109	US 1961-119830	19610627
			US	19610627

PRIORITY APPLN. INFO.:

AB A solution of 40 g. p-H2NC6H4COMe in 250 ml. H2O and 143 ml. concentrated HCl was

added at 0-5° dropwise and with stirring to a solution of 21 g. NaNO2 in 200 ml. H2O. To the resulting solution was added 60.3 g. ethyl  $\alpha$ -(2-dimethylaminoethyl)acetoacetate and 63 g. NaOAc. The mixture was maintained at pH 6-7 with 3N NaOH, stirred in the cold 2 hrs., basified, and extracted with CHCl3 to yield 65 g. ethyl  $\alpha$ -oxo- $\gamma$ -dimethylaminobutyrate p-acetylphenylhydrazones (I), m. 78-80° (petr.-ether). A mixture of 43 g. I and 430 g. polyphosphoric acid was heated with stirring at 100-10° for 2 hrs., poured into ice-H2O, basified with 3N NaOH, and extracted with 400 ml. CHCl3. The residue from the CHCl3 solution was dissolved in EtOAc and treated with ethereal HCl to yield 17.8 g. 5-acetyl-2-carbethoxygramine-HCl, m. 211-14° (MeCN). These comds. lower blood pressure, increase coronary flow, exhibit anti-serotonin activity, and depress the central nervous system.

L8 ANSWER 9 OF 9 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN

ACCESSION NUMBER: 1949:21300 BIOSIS

DOCUMENT NUMBER: PREV19492300021501; BA23:21501



TITLE: Glutinosin. A fungistatic metabolic product of the mould  
Metarrhizium glutinosum S. Pope.  
AUTHOR(S): BRIAN, P. W.; CURTIS, P. J.; HEMMING, H. G.  
CORPORATE SOURCE: Butterwick Re. Labs., Welwyn, Herts, Eng.  
SOURCE: PROC ROY SOC SER B BIOL SCI, (1947) Vol. 135, No. 878, pp.  
106-132.  
DOCUMENT TYPE: Article  
FILE SEGMENT: BA  
LANGUAGE: Unavailable  
ENTRY DATE: Entered STN: May 2007  
Last Updated on STN: May 2007

AB When *M. glutinosum* is grown at pH 3.6 on Raulin-Thom medium, the glutinosin content reaches a peak of 256 B.A. U./ml. as measured by the inhibition of germination of *Botrytis allii* conidia. Bacteriostatic action against *Staphylococcus aureus* or *Salmonella typhi* is minimal. Biotin and aneurin supplements produce marked stimulation of sporulation and slight increase in glutinosin production. Nitrate stimulates growth but not glutinosin production. Inorganic ammonium salts are ineffective for growth or glutinosin production, but peptone or the addition of 0.1-1% of tartaric, malic, succinic, malonic, oxalic, citric, acetic, glycollic, pyruvic, aspartic or glutamic acids to a medium containing 0.16% (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub>, 5% dextrose, and minerals, is highly effective in promoting growth and glutinosin production, pointing to a need for intermediates of the Krebs cycle for effective utilization of carbohydrate when the ammonia path of assimilation is followed. Cultures with the monocarboxylic acids were at initial pH 6.5 to lessen the toxicity of the free acid, but cultures with the dicarboxylic acids were started at pH 4. Glutinosin was extracted from culture medium by petroleum ether to yield on evaporation a gummy crystalline mass, which recrystallized from ethanol as a white microcrystalline solid, glutinosin, C<sub>48</sub>H<sub>60</sub>O<sub>16</sub>. For routine production, a solution containing 50 g. glucose, 0.75 g. H<sub>3</sub>PO<sub>4</sub>, 10 g. malic acid, 0.5 g. MgSO<sub>4</sub>, and 1 ml. minor element concentrate in 1 liter adjusted to pH 4 with a solution that is 2.5 N with respect to both KOH and NH<sub>4</sub>OH, yields 100 mg. glutinosin when the glutinosin is absorbed on 10 g. charcoal and extracted 6 hrs. with benzene in a Soxhlet extractor. Aqueous solns. of glutinosin are relatively stable at pH 3, with 50% activity loss in 9 days at 25[degree]C or on autoclaving 20 min. at 15 lbs. Toxicity data are given for 34 fungi and 10 bacteria. *M. glutinosum* cultures produce a volatile, water-soluble dermatitic substance, so that ventilation and barrier creams must be used when working with culture filtrates. ABSTRACT AUTHORS: E. H. Shaw, Jr

=> s trifunctional  
L9 5853 TRIFUNCTIONAL

=> s L1 and L9  
L10 88 L1 AND L9

=> s aspartyl  
L11 18645 ASPARTYL

=> s L10 and L11  
L12 1 L10 AND L11

=> d L12 ibib abs

L12 ANSWER 1 OF 1 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN  
ACCESSION NUMBER: 2002:583437 BIOSIS  
DOCUMENT NUMBER: PREV200200583437  
TITLE: Trifunctional conjugation reagents. Reagents that

contain a biotin and a radiometal chelation moiety for application to extracorporeal affinity adsorption of radiolabeled antibodies.

AUTHOR(S): Wilbur, D. Scott [Reprint author]; Chyan, Ming-Kuan; Hamlin, Donald K.; Kegley, Brian B.; Nilsson, Rune; Sandberg, Bengt E. B.; Brechbiel, Martin

CORPORATE SOURCE: Department of Radiation Oncology, University of Washington, 2121 N. 35th Street, Seattle, WA, 98103-9103, USA  
dswilbur@u.washington.edu

SOURCE: Bioconjugate Chemistry, (September-October, 2002) Vol. 13, No. 5, pp. 1079-1092. print.  
CODEN: BCCHE5. ISSN: 1043-1802.

DOCUMENT TYPE: Article

LANGUAGE: English

ENTRY DATE: Entered STN: 13 Nov 2002  
Last Updated on STN: 13 Nov 2002

AB A method of removing radiolabeled monoclonal antibodies (mAbs) from blood using a device external to the body, termed extracorporeal affinity-adsorption (EAA), is being evaluated as a means of decreasing irradiation of noncancerous tissues in therapy protocols. The EAA device uses an avidin column to capture biotinylated-radiolabeled mAbs from circulated blood. In this investigation, three trifunctional reagents have been developed to minimize the potential deleterious effect on antigen binding brought about by the combination of radiolabeling and biotinylation of mAbs required in the EAA approach. The studies focused on radiolabeling with <sup>111</sup>In and <sup>90</sup>Y, so the chelates CHX-A"-DTPA and DOTA, which form stable attachments to these radionuclides, were incorporated in the trifunctional reagents. The first trifunctional reagent prepared did not incorporate a group to block the biotin cleaving enzyme biotinidase, but the two subsequent reagents coupled aspartic acid to the biotin carboxylate for that purpose. All three reagents used 4,7,10-trioxo-1,13-tridecanediamine as water-soluble spacers between an aminoisophthalate core and the biotin or chelation group. The mAb conjugates were radioiodinated to evaluate cell binding as a function of substitution. Radioiodination was used so that a direct comparison with unmodified mAb could be made. Evaluation of the number of conjugates per antibody versus cell binding immunoreactivities indicated that minimizing the number of conjugates was best. Interestingly, a decrease of radioiodination yield as a function of the number of isothiocyanate containing conjugates per mAb was noted. The decreased yields were presumably due to the presence of thiourea functionality formed in the conjugation reaction. Radiolabeling with <sup>111</sup>In and <sup>90</sup>Y was facile at room temperature for conjugates containing the CHX-A", but elevated temperature (e.g., 45degreeC) was required to obtain good yields with the DOTA chelate. Stability of <sup>90</sup>Y labeled mAb in serum, and when challenged with 10 mM EDTA, was high. However, challenging the <sup>90</sup>Y labeled mAb with 10 mM DTPA demonstrated high stability for the DOTA containing conjugate, but low stability for the CHX-A" containing conjugate. Thus, the choice between these two chelating moieties might be made on requirements for facile and gentle labeling versus very high in vivo stability. Application of the trifunctional biotinylation reagents to the blood clearance of labeled antibodies in EAA is under investigation. The new reagents may also be useful for other applications.

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LOGOFF? (Y)/N/HOLD:y

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

55.42

55.63

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-6.40	-6.40

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